**Example 10**

**siRNA constructs prepared and tested against eIF4E and Survivin targets**

Selected siRNA constructs were prepared and tested for their ability to lower targeted RNA as measured by RT-PCR. The $IC_{50}$ of each construct was determined.

| SEQ ID No: | ISIS No: | Constructs targeted to eIF4E (5'-3'): |
|---|---|---|
| | | **Mixed 2'-F/2'-$OCH_3$ as/s RNA** |
| 25 (as) eIF4E | 349890 | UfGfUfCfAfUmAmUfUfCmCmUfGfGfAfUfCfCmUmUm |
| 26 (s) eIF4E | 338935 | AAGGAUCCAGGAAUAUGACA |
| 27 (as) eIF4E | 349891 | UfCfCfUfGfGmAmUfCfCmUmUfCfAfCfCfAfAmUmGm |
| 28 (s) eIF4E | 338939 | CAUUGGUGAAGGAUCCAGGA |
| 29(as) eIF4E | 349892 | UfCfUfUfAfUmCmAfCfCmUmUfUfAfGfCfUfCmUmAm |
| 30 (s) eIF4E | 338943 | UAGAGCUAAAGGUGAUAAGA |
| 31 (as) eIF4E | 351097 | AfUfAfCfUfCmAmGfAfAmGmGfUfGfUfCfUfUmCmUm |
| 32 (s) eIF4E | 338952 | AGAAGACACCUUCUGAGUAU |
| | | **Full P=S RNA as/s full 2'-$OCH_3$** |
| 35 (as) Survivin | 346280 | UrsUrsUrsGrsArsArsArsArsUrsGrsUrsUrsGrsArsUrsCrsUrsCrsCrsU |
| 34 (s) Survivin | 352512 | GmGmAmGmAmUmCmAmAmCmAmUmUmUmUmCmAmAmAm |
| | | **Full P=S RNA as/s full 2'-$OCH_3$ except terminal nucleosides** |
| 35 (as) Survivin | 346280 | UrsUrsUrsGrsArsArsArsArsUrsGrsUrsUrsGrsArsUrsCrsUrsCrsCrsU |
| 34 (s) Survivin | 352513 | GGmAmGmAmUmCmAmAmCmAmUmUmUmUmCmAmAmA |

| SEQ ID No: | ISIS No: | Constructs targeted to Survivin (5'-3'): |
|---|---|---|
| | | **Mixed 2'-F/2'-$OCH_3$ as/s RNA** |
| 33 (as) Survivin | 355711 | UfUfUfGfAfAmAmAfUfGmUmUfGfAfUfCfUmCmCm |
| 34 (s) Survivin | 343868 | GGAGAUCAACAUUUUCAAA |

Lowercase f indicates that the preceding nucleoside is a 2'-F nucleoside (Cf =2'-F cytidine). Lowercase m indicates the previous nucleoside is a 2'-$OCH_3$ nucleoside. Lowercase s (second lowercase letter following a U, A, G, or C) after a lowercase letter indicates that the internucleoside linkage is a phosphorothioate internucleoside linkage (rs = the sugar is ribose and the internucleoside linkage is phosphorothioate.

The above constructs were tested in HeLa cells, MH-S cells or U-87 MG cells using a standard assay as shown in Example 15. The $IC_{50}$'s were calculated as shown below.